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	OTHER PUBLICATIONS (including Author, Title, Date, Pertinent Pages, Etc.)									
HAR		Ward et al., An Energy transfer protein in coelenterate bioluminescence. Characterization of the Renilla green fluorescent protein. J. Biol. Chem. 254 no 3 (1979) 781-788								
	D2	Geoghegan et al., Site directed double fluorescent tagging of human renin and collagenase (MMP-1) substrate peptides using the periodate oxidation of N-terminal serine. An apparently general strategy for provision of energy transfer substrates for proteases. Bioconjugate Chemistry 4 no 6, (1993) 537-544								
	D3	Wu et al. Resonance energy transfer: Methods and applications. Analytical Biochemistry 218, 1194, p1-13 Academic press Inc., Duluth, NM								
	D4	Inouye and Tsuji, Expression of the gene and fluorescence characteristics of the recombinant proteins. FEBS Letters 341:277-280 (1994)								
	D5	Mitra et al., Fluorescence resonance energy transfer between blue emitting and red-shifted excitation derivatives of the green fluorescent protein. Gene, 173:13-17 (1996)								

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		Delagrave et al., "Ro (1995)	ed-Shifted Ex	citation Mutants of	the Green Fluorescent Pro	tein", <u>Bi</u>	o/ recnnoto	97, 13:13	124			
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			9(4): pp. 650-655 (1995) ubitt et al., "Understanding, improving and using green fluorescent proteins, <u>TIBS</u> , (1995) 20:448-455									
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· v paritimation of the control of t	BM	Engineering, 1991, 4	:457-461		e of an exported protein w				
					ictoria green-fluorescent			2, 111:22	9-233
					lling," <u>Trends Cell Biol</u> ,				
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